SEP 1 6 2008

Sequence Listing

Kelley, Robert F. Hymowitz, Sarah Lindstrom, Stephanie Ho

- <120> APO-2 LIGAND/TRAIL VARIANTS AND USES THEREOF
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- <140> US 10/519,647
- <141> 2006-01-10
- <150> PCT/US03/019750
- <151> 2003-06-23
- <150> US 60/391,050
- <151> 2002-06-24
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- <213> Homo sapiens
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- Val Ala Val Thr Tyr Val Tyr Phe Thr Asn Glu Leu Lys Gln Met 35 40 45
- Gln Asp Lys Tyr Ser Lys Ser Gly Ile Ala Cys Phe Leu Lys Glu
 50 55 60
- Asp Asp Ser Tyr Trp Asp Pro Asn Asp Glu Glu Ser Met Asn Ser
 65 70 75
- Pro Cys Trp Gln Val Lys Trp Gln Leu Arg Gln Leu Val Arg Lys 80 85 90
- Met Ile Leu Arg Thr Ser Glu Glu Thr Ile Ser Thr Val Gln Glu 95 100 105
- Lys Gln Gln Asn Ile Ser Pro Leu Val Arg Glu Arg Gly Pro Gln
 110 115 120
- Arg Val Ala Ala His Ile Thr Gly Thr Arg Gly Arg Ser Asn Thr
 125 130 135
- Leu Ser Ser Pro Asn Ser Lys Asn Glu Lys Ala Leu Gly Arg Lys
 140 145 150
- Ile Asn Ser Trp Glu Ser Ser Arg Ser Gly His Ser Phe Leu Ser
 155 160 165
- Asn Leu His Leu Arg Asn Gly Glu Leu Val Ile His Glu Lys Gly 170 175 180

Phe Tyr Tyr Ile Tyr Ser Gln Thr Tyr Phe Arg Phe Gln Glu Glu 190 Ile Lys Glu Asn Thr Lys Asn Asp Lys Gln Met Val Gln Tyr Ile 200 205 Tyr Lys Tyr Thr Ser Tyr Pro Asp Pro Ile Leu Leu Met Lys Ser Ala Arg Asn Ser Cys Trp Ser Lys Asp Ala Glu Tyr Gly Leu Tyr 230 235 240 Ser Ile Tyr Gln Gly Gly Ile Phe Glu Leu Lys Glu Asn Asp Arg 250 Ile Phe Val Ser Val Thr Asn Glu His Leu Ile Asp Met Asp His 260 265 Glu Ala Ser Phe Phe Gly Ala Phe Leu Val Gly

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Trp Val Ile Leu Val Val Thr Leu Val Val Pro Leu Leu Val 245 250 Ala Val Leu Ile Val Cys Cys Cys Ile Gly Ser Gly Cys Gly Gly Asp Pro Lys Cys Met Asp Arg Val Cys Phe Trp Arg Leu Gly Leu Leu Arg Gly Pro Gly Ala Glu Asp Asn Ala His Asn Glu Ile Leu 290 295 Ser Asn Ala Asp Ser Leu Ser Thr Phe Val Ser Glu Gln Met Glu Ser Gln Glu Pro Ala Asp Leu Thr Gly Val Thr Val Gln Ser 320 Pro Gly Glu Ala Gln Cys Leu Leu Gly Pro Ala Glu Ala Glu Gly 335 340 Ser Gln Arg Arg Leu Leu Val Pro Ala Asn Gly Ala Asp Pro 355 Thr Glu Thr Leu Met Leu Phe Phe Asp Lys Phe Ala Asn Ile Val Pro Phe Asp Ser Trp Asp Gln Leu Met Arg Gln Leu Asp Leu Thr 380 385 Lys Asn Glu Ile Asp Val Val Arg Ala Gly Thr Ala Gly Pro Gly Asp Ala Leu Tyr Ala Met Leu Met Lys Trp Val Asn Lys Thr Gly 415 410 Arg Asn Ala Ser Ile His Thr Leu Leu Asp Ala Leu Glu Arg Met 425 430 Glu Glu Arg His Ala Lys Glu Lys Ile Gln Asp Leu Leu Val Asp Ser Gly Lys Phe Ile Tyr Leu Glu Asp Gly Thr Gly Ser Ala Val 460

Ser Leu Glu

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<211> 411

<212> PRT

<213> Homo sapiens

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L	eu .	Ala	Pro	Gln	Gln 65	Arg	Ala	Ala	Pro	Gln 70	Gln	Lys	Arg	Ser	Ser 75					
P	ro	Ser	Glu	Gly	Leu 80	Cys	Pro	Pro	Gly	His 85	His	Ile	Ser	Glu	Asp 90					
G:	ly .	Arg	Asp	Cys	Ile 95	Ser	Cys	Lys	Tyr	Gly 100	Gln	Asp	Tyr	Ser	Thr 105					
Н	is	Trp	Asn	Asp	Leu 110	Leu	Phe	Cys	Leu	Arg 115	Cys	Thr	Arg	Cys	Asp 120					
S	er	Gly	Glu	Val	Glu 125	Leu	Ser	Pro	Cys	Thr 130	Thr	Thr	Arg	Asn	Thr 135					
V	al	Cys	Gln	Cys	Glu 140	Glu	Gly	Thr	Phe	Arg 145	Glu	Glu	Asp	Ser	Pro 150					
G)	lu :	Met	Cys	Arg	Lys 155	Cys	Arģ	Thr	Gly	Cys 160	Pro	Arg	Gly	Met	Val 165			, .		
L	ys	Val	Gly	Asp	Cys 170	Thr	Pro	Trp	Ser	Asp 175	Ile	Glu	Cys	Val	His 180	-				
L	ys	Glu	Ser	Gly	Ile 185	Ile	Ile	Gly	Val	Thr 190	Val	Ala	Ala	Val	Val 195					
L	eu	Ile	Val	Ala	Val 200	Phe	V _a l	Cys	Lys	Ser 205	Leu	Leu	Trp	Lys	Lys 210		•			
, V	al	Leu	Pro	Tyr	Leu 215	Lys	Gly	Ile	Cys	Ser 220	Gly	Gly	Gly	Gly	Asp 225					
P	ro	Glu	Arg	Val	Asp 230	_	Ser	Ser	Gln	Arg 235		Gly	Ala	Glu	Asp 240					
A	sn	Val	Leu	Asn	Glu 245	Ile	Val	Ser	Ile	Leu 250	Gln	Pro	Thr	Gln	Val 255					
					Met 260					265					270				,	
					Ser 275		_			280					285					
					Arg 290					295					300					
					Pro 305					310					315					
					Val 320					325					330					
	_				Met 335					340					345					
A.	la .	Ala	Gly	His	Arg 350	Asp	Thr	Leu	Tyr	Thr 355	Met	Leu	Ile	Lys	Trp 360					
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Val Asn Lys Thr Gly Arg Asp Ala Ser Val His Thr Leu Leu Asp 365 370 Ala Leu Glu Thr Leu Gly Glu Arg Leu Ala Lys Gln Lys Ile Glu Asp His Leu Leu Ser Ser Gly Lys Phe Met Tyr Leu Glu Gly Asn Ala Asp Ser Ala Leu Ser 410 <210> 6 <211> 440 <212> PRT <213> Homo sapiens <400> 6 Met Glu Gln Arg Gly Gln Asn Ala Pro Ala Ala Ser Gly Ala Arg Lys Arg His Gly Pro Gly Pro Arg Glu Ala Arg Gly Ala Arg Pro Gly Pro Arg Val Pro Lys Thr Leu Val Leu Val Val Ala Ala Val Leu Leu Val Ser Ala Glu Ser Ala Leu Ile Thr Gln Gln Asp Leu Ala Pro Gln Gln Arg Ala Ala Pro Gln Gln Lys Arg Ser Ser Pro Ser Glu Gly Leu Cys Pro Pro Gly His His Ile Ser Glu Asp Gly Arg Asp Cys Ile Ser Cys Lys Tyr Gly Gln Asp Tyr Ser Thr 100 95 His Trp Asn Asp Leu Leu Phe Cys Leu Arg Cys Thr Arg Cys Asp Ser Gly Glu Val Glu Leu Ser Pro Cys Thr Thr Thr Arg Asn Thr Val Cys Gln Cys Glu Glu Gly Thr Phe Arg Glu Glu Asp Ser Pro 140 Glu Met Cys Arg Lys Cys Arg Thr Gly Cys Pro Arg Gly Met Val Lys Val Gly Asp Cys Thr Pro Trp Ser Asp Ile Glu Cys Val His 170 175 Lys Glu Ser Gly Thr Lys His Ser Gly Glu Ala Pro Ala Val Glu 185 190 Glu Thr Val Thr Ser Ser Pro Gly Thr Pro Ala Ser Pro Cys Ser Leu Ser Gly Ile Ile Ile Gly Val Thr Val Ala Ala Val Val Leu 220

Ile Val Ala Val Phe Val Cys Lys Ser Leu Leu Trp Lys Lys Val 235 Leu Pro Tyr Leu Lys Gly Ile Cys Ser Gly Gly Gly Asp Pro Glu Arg Val Asp Arg Ser Ser Gln Arg Pro Gly Ala Glu Asp Asn Val Leu Asn Glu Ile Val Ser Ile Leu Gln Pro Thr Gln Val Pro 275 280 Glu Gln Glu Met Glu Val Gln Glu Pro Ala Glu Pro Thr Gly Val 295 Asn Met Leu Ser Pro Gly Glu Ser Glu His Leu Leu Glu Pro Ala 305 Glu Ala Glu Arg Ser Gln Arg Arg Leu Leu Val Pro Ala Asn 320 325 Glu Gly Asp Pro Thr Glu Thr Leu Arg Gln Cys Phe Asp Asp Phe Ala Asp Leu Val Pro Phe Asp Ser Trp Glu Pro Leu Met Arg Lys 355 Leu Gly Leu Met Asp Asn Glu Ile Lys Val Ala Lys Ala Glu Ala 370 365 Ala Gly His Arg Asp Thr Leu Tyr Thr Met Leu Ile Lys Trp Val Asn Lys Thr Gly Arg Asp Ala Ser Val His Thr Leu Leu Asp Ala 400 Leu Glu Thr Leu Gly Glu Arg Leu Ala Lys Gln Lys Ile Glu Asp 410 415 420 His Leu Leu Ser Ser Gly Lys Phe Met Tyr Leu Glu Gly Asn Ala Asp Ser Ala Met Ser <210> 7 <211> 161 <212> PRT <213> Homo sapiens <400>7Arg Val Ala Ala His Ile Thr Gly Thr Arg Gly Arg Ser Asn Thr Leu Ser Ser Pro Asn Ser Lys Asn Glu Lys Ala Leu Gly Arg Lys Ile Asn Ser Trp Glu Ser Ser Arg Ser Gly His Ser Phe Leu Ser Asn Leu His Leu Arg Asn Gly Glu Leu Val Ile His Glu Lys Gly

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 Ile Lys Glu Asn Thr Lys Asn Asp Lys Gln Met Val Gln Tyr Ile
Tyr Lys Tyr Thr Ser Tyr Pro Asp Pro Ile Leu Leu Met Lys Ser
Ala Arg Asn Ser Cys Trp Ser Lys Asp Ala Glu Tyr Gly Leu Tyr
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                                     115
Ser Ile Tyr Gln Gly Gly Ile Phe Glu Leu Lys Glu Asn Asp Arg
Ile Phe Val Ser Val Thr Asn Glu His Leu Ile Asp Met Asp His
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                                     145
Glu Ala Ser Phe Phe Gly Ala Phe Leu Val Gly
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Tyr Phe Val Tyr Ser Gln Val Val Phe Ser Gly Lys Ala Tyr Ser
Pro Lys Ala Thr Ser Ser Pro Leu Tyr Leu Ala His Glu Val Gln
Leu Phe Ser Ser Gln Tyr Pro Phe His Val Pro Leu Leu Ser Ser
Gln Lys Met Val Tyr Pro Gly Leu Gln Glu Pro Trp Leu His Ser
                  95
                                     100
                                                          105
Met Tyr His Gly Ala Ala Phe Gln Leu Thr Gln Gly Asp Gln Leu
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Ser Thr His Thr Asp Gly Ile Pro His Leu Val Leu Ser Pro Ser
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Thr Val Phe Phe Gly Ala Phe Ala Leu
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130

125

Phe Gly Leu Leu Lys Leu 140 <210> 11 <211> 137 <212> PRT

Tyr Lys

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130

His Thr Leu Met Lys Gly Gly Ser Thr Lys Tyr Trp Ser Gly Asn 95 100 105

Ser Glu Phe His Phe Tyr Ser Île Asn Val Gly Gly Phe Phe Lys

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Leu Leu Asp Pro Asp Gln Asp Ala Thr Tyr Phe Gly Ala Phe Lys
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Val Arg

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